

## **Amendments to the Claims**

This listing of claims will replace all prior versions, and listings, of claims in the specification:

### **Listing of Claims:**

1. (original) The use of an isolated proton-sensing GPCR polypeptide in the development of a medicament for disease and medical conditions in which proton homeostasis is imbalanced.
2. (original) The use of an isolated proton-sensing GPCR polypeptide in the development of a medicament for disease and medical conditions in which proton homeostasis is imbalanced; said polypeptide is selected from one of the groups consisting of:
  - (a) an isolated polypeptide encoded by a polynucleotide comprising the polynucleotide sequence of human OGR1 (accession number: NM\_003485.1), rat OGR1 (accession number: XM\_234483), mouse OGR1 (accession number: NM\_175493), bovine OGR1 (accession number: NM\_174329), preferably human OGR1 (accession number: NM\_003485.1), human GPM (accession number: NM\_005282), mouse GPR4 (accession number: NM\_175668), human TDAG8 (accession number: NM\_003608) and mouse TDAG8 (accession number: NM\_008152);
  - (b) an isolated proton sensing GPCR polypeptide comprising a polypeptide sequence having at least 20% to the polypeptide sequence of SEQ ID NO: 1;
  - (c) an isolated proton sensing GPCR polypeptide comprising a polypeptide sequence having at least 20% to the polypeptide sequence of SEQ ID NO: 3;
  - (d) an isolated proton sensing GPCR polypeptide comprising a polypeptide sequence having at least 20% identity to the polypeptide sequence of SEQ ID NO: 4;
  - (e) an isolated polypeptide comprising the polypeptide sequence of SEQ ID NO: 1, SEQ ID NO: 3 or SEQ ID NO: 4;
  - (f) an isolated proton sensing GPCR polypeptide having at least 20% identity to the polypeptide sequence of SEQ ID NO: 1;
  - (g) an isolated proton sensing GPCR polypeptide having at least 20% identity to the polypeptide sequence of SEQ ID NO: 3;
  - (h) an isolated proton sensing GPCR polypeptide having at least 20% identity to the polypeptide sequence of SEQ ID NO: 4;
  - (i) the polypeptide sequences of SEQ ID NO: 1, SEQ ID NO: 3, or SEQ ID NO: 4; and
  - (j) polypeptides in (a) to (i) which show a pH dependent Inositol phosphate formation in CCL39 hamster fibroblast cells or a pH dependent signal in the cAMP luciferase reporter assay in CHOK1 CRE-luc cells or CCL39 CRE-luc cells.

3. (currently amended) The use of the isolated polypeptide as claimed in claim 1 ~~or~~ 2, said polypeptide comprising the polypeptide sequence of SEQ ID NO: 1, SEQ ID NO: 3 or SEQ ID NO: 4.
4. (currently amended) The use of the isolated polypeptide as claimed in claim 1 ~~or~~ 2, said polypeptide consisting of the polypeptide sequence of SEQ ID NO: 1, SEQ ID NO: 3 or SEQ ID NO: 4.
5. (original) The use of an isolated polynucleotide in the development of a medicament for the prevention and treatment of diseases and medical conditions in which proton homeostasis is imbalanced; said polynucleotide is selected from one of the groups consisting of:
  - (a) an isolated polynucleotide comprising the polynucleotide sequence of human OGR1 (accession number: NM\_003485.1), rat OGR1 (accession number: XM\_234483), mouse OGR1 (accession number: NM\_175493), bovine OGR1 (accession number: NM\_174329), preferably human OGR1 (accession number: NM\_003485.1), human GPR4 (accession number: NM\_005282), mouse GPR4 (accession number: NM\_175668), human TDAG8 (accession number: NM\_003608) and mouse TDAG8 (accession number: NM\_008152);
  - (b) an isolated polynucleotide encoding a proton sensing GPCR polypeptide sequence having at least 20% identity to the polypeptide sequence of SEQ ID NO: 1;
  - (c) an isolated polynucleotide encoding a proton sensing GPRC polypeptide sequence having at least 20% identity to the polypeptide sequence of SEQ ID NO: 3;
  - (d) an isolated polynucleotide encoding a proton sensing GPRC polypeptide sequence having at least 20% identity to the polypeptide sequence of SEQ ID NO: 4;
  - (f) an isolated polynucleotide comprising the polynucleotide sequence of human OGR1 (accession number: NM\_003485.1), rat OGR1 (accession number: XM\_234483), mouse OGR1 (accession number: NM\_175493), bovine OGR1 (accession number: NM\_174329), preferably human OGR1 (accession number: NM\_003485.1), human GPR4 (accession number: NM\_005282), mouse GPR4 (accession number: NM\_175668), human TDAG8 (accession number: NM\_003608) and mouse TDAG8 (accession number: NM\_008152);
  - (g) the polynucleotide sequences of human OGR1 (accession number: NM\_003485.1), rat OGR1 (accession number: XM\_234483), mouse OGR1 (accession number: NM\_175493), bovine OGR1 (accession number: NM\_174329), preferably human OGR1 (accession number: NM\_003485.1), human GPR4 (accession number: NM\_005282), mouse GPR4 (accession number: NM\_175668), human TDAG8 (accession number: NM\_003608) and mouse TDAG8 (accession number: NM\_008152); and
  - (h) polynucleotides in (a) to (g) which encode for a polypeptide that show a phi dependent Inositol phosphate formation in CCL39 hamster fibroblast cells or a pH dependent signal in the cAMP luciferase reporter assay in CHOK1 CRE-luc cells or CCL39 CRE-luc cells.

6. (currently amended) The use of an antibody, which specifically binds to a polypeptide of ~~any one of the claims 1 to 4~~, for the manufacture of a medicament for the prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalanced; a pharmaceutical composition comprising an antibody for the prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalances, said antibody specifically binds to a polypeptide of ~~any one of the claims 1 to 4~~; or a method of prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalanced comprising administering to a subject in need of such prevention and/or treatment an effective amount of an antibody, said antibody specifically binds to a polypeptide of ~~any one of the claims 1 to 4~~.

7. (currently amended) A method for screening to identify compounds that antagonize the proton-sensing activity of the polypeptides according to ~~any one of claims 1 to 4~~.

8. (currently amended) A method for screening to identify compounds that agonize the proton-sensing activity of the polypeptides according to ~~any one of claims 1 to 4~~.

9. (currently amended) A method for screening to identify compounds that stimulate or inhibit the function or expression level of the polypeptides of ~~claims 1 to 4~~ comprising a method selected from the group consisting of:

(a) measuring or, detecting, quantitatively or qualitatively, the binding of a candidate compound to the polypeptide (or to the cells or membranes expressing the polypeptide) or a fusion protein thereof by means of a label directly or indirectly associated with the candidate compound;

(b) measuring the competition of binding of a candidate compound to the polypeptide (or to the cells or membranes expressing the polypeptide) or a fusion protein thereof in the presence of a labelled competitor;

(c) testing whether the candidate compound results in a signal generated by activation or inhibition of the polypeptide, using detection systems appropriate to the cells or cell membranes expressing the polypeptide; or

(d) detecting the effect of a candidate compound on the production of mRNA encoding said polypeptide or said polypeptide in cells, using for instance, an ELISA assay.

10. (original) A method of prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalanced comprising administering to a subject in need of such prevention and/or treatment an effective amount of an antagonist obtainable from the method of claim 7.

11. (original) A method of prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalanced comprising administering to a subject in need of such

prevention and/or treatment an effective amount of an agonist obtainable from the method of claim 8.

12. (original) A pharmaceutical composition for the prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalanced comprising an antagonist obtainable from the method of claim 7.

13. (original) A pharmaceutical composition for the prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalanced comprising an agonist obtainable from the method of claim 8.

14. (currently amended) A diagnostic kit comprising an antibody against a polypeptide according to ~~any one of claims 1 to 4~~.

15. (currently amended) A diagnostic kit comprising a pharmaceutical preparation for the prevention and/or treatment of diseases and medical conditions in which proton homeostasis is imbalances, said pharmaceutical preparation comprising an antibody against a polypeptide according to ~~any one of claims 1 to 4~~.